

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Applicant: Huiping GUAN et al.  
Title: 3-(4-AMIDOPYRROL-2-YLMETHYLIDENE)-2-INDOLINONE  
DERIVATIVES AS PROTEIN KINASE INHIBITORS  
Prior Appl. No.: 10/076,140  
Prior Appl. Filing Date: 02/15/2002  
Examiner: Unassigned  
Art Unit: Unassigned

**INFORMATION DISCLOSURE STATEMENT**  
**UNDER 37 CFR §1.56**

Mail Stop PATENT APPLICATION  
Commissioner for Patents  
PO Box 1450  
Alexandria, Virginia 22313-1450

Sir:

Applicants submit herewith on Form PTO/SB/08 a listing of the documents cited by or submitted to the U.S. PTO in parent application Serial No. 10/076,140, filed 02/15/2002. As provided in 37 CFR §1.98(d), copies of the documents are not being provided since they were previously submitted to the United States Patent & Trademark Office in the above-identified parent application.

The submission of any document herewith, which is not a statutory bar, is not intended as an admission that such document constitutes prior art against the claims of the present application or that such document is considered material to patentability as defined in 37 CFR §1.56(b). Applicants do not waive any rights to take any action which would be appropriate to antedate or otherwise remove as a competent reference any document which is determined to be a *prima facie* art reference against the claims of the present application.

**TIMING OF THE DISCLOSURE**

The listed documents are being submitted in compliance with 37 CFR §1.97(b), within three (3) months of the filing date of the application.

**RELEVANCE OF EACH DOCUMENT**

Applicants respectfully request that the listed documents be considered by the Examiner and be made of record in the present application and that an initialed copy of Form PTO/SB/08 be returned in accordance with MPEP §609.

The Commissioner is hereby authorized to charge any additional fees which may be required regarding this application under 37 CFR §§ 1.16-1.17, or credit any overpayment, to Deposit Account No. 19-0741. Should no proper payment be enclosed herewith, as by a check being in the wrong amount, unsigned, post-dated, otherwise improper or informal or even entirely missing, the Commissioner is authorized to charge the unpaid amount to Deposit Account No. 19-0741.

Date

9/8/03

FOLEY & LARDNER  
Customer Number: 22428



22428

PATENT TRADEMARK OFFICE

Telephone: (202) 672-5475  
Facsimile: (202) 672-5399

Respectfully submitted,

By

Beth A. Burrous  
Attorney for Applicant  
Registration No. 35,087

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First Named Inventor	Huiping GUAN et al.
Group Art Unit	Unassigned
Examiner Name	Unassigned
Attorney Docket Number	034536-0188

Sheet 1 of 30

**U.S. PATENT DOCUMENTS**

Examiner Initials*	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number	Kind Code <sup>2</sup> (if known)			
	B1	Re. 35,096	E	Taniguchi et al.	11-21-1995	
	B2	Re. 36,256	E	Spada et al.	07-20-1999	
	B3	2,622,980		Copeland	12-23-1952	
	B4	2,872,372		Hull	02-03-1959	
	B5	2,968,557		Burgandt et al.	01-17-1961	
	B6	3,140,180		Fritz	07-07-1964	
	B7	3,308,134		Plotneiks	03-07-1967	
	B8	3,551,571		Pachter et al.	12-29-1970	
	B9	3,564,016		Schoen et al.	02-16-1971	
	B10	3,715,364		Hoff	02-06-1973	
	B11	3,880,871		Haugwitz et al.	04-29-1975	
	B12	3,922,163		Church et al.	11-25-1975	
	B13	4,002,643		Carson	01-11-1977	
	B14	4,002,749		Rovnyak	01-11-1977	
	B15	4,053,613		Rovnyak et al.	10-11-1977	
	B16	4,070,366		Gregorovich et al.	01-24-1978	
	B17	4,259,345		Cross et al.	03-31-1981	
	B18	4,343,923		Lenox et al.	08-10-1982	

Examiner  
SignatureDate  
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	B19	4,376,110		David et al.	03-08-1983	
	B20	4,436,892		Zondler et al.	03-13-1984	
	B21	4,489,089		Wright, Jr. et al.	12-18-1984	
	B22	4,493,842		Furazawa et al.	01-15-1985	
	B23	4,560,700		Schnettler et al.	12-24-1985	
	B24	4,628,105		Schmid et al.	12-09-1986	
	B25	4,642,309		Michel et al.	02-10-1987	
	B26	4,678,798		Rentzea et al.	07-07-1987	
	B27	4,826,847		Michel et al.	05-02-1989	
	B28	4,853,403		Shiraishi et al.	08-01-1989	
	B29	4,853,404		Takamura et al.	08-01-1989	
	B30	4,868,304		Larock	09-19-1989	
	B31	4,924,000		Rentzea et al.	05-08-1990	
	B32	4,966,849		Vallee et al.	10-30-1990	
	B33	4,971,996		Shiraishi et al.	11-20-1990	
	B34	4,987,146		Rohde et al.	01-22-1991	
	B35	5,043,348		Zoller et al.	08-27-1991	
	B36	5,043,454		Wriede et al.	08-27-1991	

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		Number	Kind Code <sup>2</sup> (if known)			
	B37	5,047,554		Ehrgott et al.	09-10-1991	
	B38	5,051,417		Nadler et al.	09-24-1991	
	B39	5,057,538		Shiraishi et al.	10-15-1991	
	B40	5,082,856	A	Taniguchi et al.	01-21-1992	
	B41	5,089,516	A	Shiraishi et al.	02-18-1992	
	B42	5,124,347	A	Connor et al.	06-23-1992	
	B43	5,145,983	A	West	09-08-1992	
	B44	5,153,217	A	Taniguchi et al.	10-06-1992	
	B45	5,196,446	A	Levitzi et al.	03-23-1993	
	B46	5,202,341	A	Shiraishi et al.	04-13-1993	
	B47	5,206,261	A	Kawaguchi et al.	04-27-1993	
	B48	5,217,999	A	Levitzi et al.	06-08-1993	
	B49	5,258,357	A	Muenster et al.	11-02-1993	
	B50	5,278,184	A	Artico et al.	01-11-1994	
	B51	5,290,947	A	Zoller et al.	03-01-1994	
	B52	5,302,606	A	Spada et al.	04-12-1994	
	B53	5,322,950	A	Sircar et al.	06-21-1994	
	B54	5,330,992	A	Eissenstat et al.	07-19-1994	

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				<b>First Named Inventor</b>	Huiping GUAN et al.
				<b>Group Art Unit</b>	Unassigned
				<b>Examiner Name</b>	Unassigned
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		Number	Kind Code <sup>2</sup> (if known)			
	B55	5,332,736	A	Carmosin et al.	07-26-1994	
	B56	5,374,652	A	Buzzetti et al.	12-20-1994	
	B57	5,382,593	A	Le Baut et al.	01-17-1995	
	B58	5,389,661	A	Sircar et al.	02-14-1995	
	B59	5,397,787	A	Buzzetti et al.	03-14-1995	
	B60	5,409,930	A	Spada et al.	04-25-1995	
	B61	5,409,949	A	Buzzetti et al.	04-25-1995	
	B62	5,463,052	A	Haga et al.	10-31-1995	
	B63	5,565,324	A	Still et al.	10-15-1996	
	B64	5,610,173	A	Schwartz et al.	03-11-1997	
	B65	5,723,665	A	Kato et al.	03-03-1998	
	B66	5,786,488	A	Tang et al.	07-28-1998	
	B67	5,792,783	A	Tang et al.	08-11-1998	
	B68	5,834,504	A	Tang et al.	11-10-1998	
	B69	5,849,710	A	Battistini et al.	12-15-1998	
	B70	5,880,141	A	Tang et al.	03-09-1999	
	B71	5,883,113	A	Tang et al.	03-16-1999	
	B72	5,883,116	A	Tang et al.	03-16-1999	

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		Number	Kind Code <sup>2</sup> (if known)			
	B73	5,886,020	A	Tang et al.	03-23-1999	
	B74	6,130,239	A	Chen et al.	10-10-2000	
	B75	6,133,305	A	Tang et al.	10-17-2000	
	B76	6,284,894	B1	Phillion et al.	09-04-2001	
	B77	6,310,217	B1	Lehr	10-30-2001	
	B78	6,395,736		Parks et al.	05-28-2002	
	B79	6,451,838		Moon et al.	09-17-2002	
	B80	6,462,072		Hamilton et al.	10-08-2002	

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	B81	WO	88/07035	A1	KANEGAFUCHI KAGAKU KOGYO KABUSHIKI KAISHA	09-22-1988		
	B82	WO	91/13055	A2	FARMITALIA CARLO ERBA SRL	09-05-1991		
	B83	WO	91/15495	A1	PFIZER INC.	10-17-1991		
	B84	WO	92/03736	A1	SEIKAGAKU KOGYO KABUSHIKI KAISHA	03-05-1992		
	B85	WO	92/07830	A2	PFIZER INC.	05-14-1992		
	B86	WO	92/20642	A1	RHONEPOULENC RORER INTERNATIONAL	11-26-1992		
	B87	WO	92/21660	A1	PFIZER INC.	12-10-1992		

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	B88	WO	93/01182	A1	FARMITALIA CARLO ERA SRL	01-21-1993		
	B89	WO	93/23040	A1	MERCK & CO., INC.	11-25-1993		
	B90	WO	94/03427	A1	WARNER-LAMBERT COMPANY	02-17-1994		
	B91	WO	94/10202	A1	GENENTECH, INC.	05-11-1994		
	B92	WO	94/14808	A1	FARMITALIA CARLO ERBA SRL	07-07-1994		
	B93	WO	95/01349	A1	FARMITALIA CARLO ERBA SRL	01-12-1995		
	B94	WO	95/14667	A1	PFIZER INC.	06-01-1995		
	B95	WO	95/17181	A1	PHARMACIA S.P.A.	06-29-1995		
	B96	WO	95/24190	A2	SUGEN, INC.	09-14-1995		
	B97	WO	96/00226	A1	PHARMACIA S.P.A.	01-04-1996		
	B98	WO	96/16964	A1	PHARMACIA S.P.A.	06-06-1996		
	B99	WO	96/22976	A1	PHARMACIA S.P.A.	08-01-1996		
	B100	WO	96/32380	A1	PHARMACIA S.P.A.	10-17-1996		
	B101	WO	96/40116	A1	SUGEN, INC.	12-19-1996		
	B102	WO	97/25986	A1	TAIHO PHARMACEUTICAL CO., LTD.	07-24-1997		
	B103	WO	97/34920	A1	SUGEN, INC.	09-25-1997		
	B104	WO	97/36867	A1	PFIZER, INC.	10-09-1997		
	B105	WO	98/07695	A1	SUGEN, INC.	02-26-1998		

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	B106	WO	98/07835	A2	SUGEN, INC.	02-26-1998		
	B107	WO	98/24432	A2	SUGEN, INC.	06-11-1998		
	B108	WO	98/38984	A2	SUGEN, INC.	09-11-1998		
	B109	WO	98/45708	A1	SUGEN, INC.	10-15-1998		
	B110	WO	98/50356	A1	SUGEN, INC.	11-12-1998		
	B111	WO	98/56376	A1	SUGEN, INC.	12-17-1998		
	B112	WO	99/10325	A1	GLAXO GROUP LIMITED	03-04-1999		
	B113	WO	99/19325	A1	SYNTHELABO	04-22-1999		
	B114	WO	99/48868	A2	SUGEN, INC.	09-30-1999		
	B115	WO	99/52869	A1	BOEHRINGER INGELHEIM PHARMA KG	10-21-1999		
	B116	WO	99/65869	A1	BAYER AKTIENGESELLSCHAFT	12-23-1999		
	B117	WO	99/61422	A1	SUGEN, INC.	12-02-1999		
	B118	WO	00/35920	A2	F. HOFFMANN-LA ROCHE AG	06-22-2000		
	B119	WO	00/38519	A1	SUGEN, INC.	07-06-2000		
	B120	WO.	00/08202	A2	SUGEN, INC.	02-17-2000		
	B121	WO	00/56709	A1	SUGEN, INC.	09-28-2000		
	B122	WO	01/60814	A2	SUGEN, INC.	08-23-2001		
	B123	WO	01/90068	A2	SUGEN, INC. et al.	11-29-2001		
	B124	DE	878,539		Von FREYBERG, et al.	06-05-1953		X

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Prior Application Number	10/076,140
Prior Appl. Filing Date	February 15, 2002
First Named Inventor	Huiping GUAN et al.
Group Art Unit	Unassigned
Examiner Name	Unassigned
Attorney Docket Number	034536-0188

Sheet 8 of 30

## FOREIGN PATENT DOCUMENTS

Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Documents	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>6</sup>
		Office <sup>3</sup>	Number	Kind Code <sup>5</sup> (if known)				
	B125	DE	2,159,360	A	BAYER AG	06-14-1973		X
	B126	DE	2,159,361	A	BAYER AG	06-14-1973		X
	B127	DE	2,159,362		BAYER AG	06-14-1973		
	B128	DE	2,159,363	A	BAYER AG	06-14-1973		X
	B129	DE	2,321,656	A	COLGATE-PALMOLIVE CO.	11-15-1973		X
	B130	DE	3,426,419	A	BOEHRINGER MANNHEIM GMBH	01-23-1986		X
	B131	EP	0 252 713	B1	PFIZER INC.	01-13-1988		
	B132	EP	0 304 493	B1	KANEGAFUCHI KAGAKU KOGTO KABUSHIKI KAISHA	03-01-1989		
	B133	EP	0 351 213	A2	LES LABORATOIRES BEECHAM S.A.	01-17-1990		
	B134	EP	0 525 472	A2	FARMITALIA CARLO ERBA SRL	02-03-1993		
	B135	EP	0 566 226	B1	ZENECA LIMITED	10-20-1993		
	B136	EP	0 580 502	B1	ADIR ET COMPAGNIE	01-26-1994		X
	B137	EP	0 626 377	B1	SHIONOGI & CO., LTD.	11-30-1994		
	B138	EP	0 632 102	A1	BAYER AG	01-04-1995		X
	B139	EP	0 662 473	A1	PHARMACIA S.P.A.	07-12-1995		
	B140	EP	0 788 890	A1	AGFA-GEVAERT	08-13-1997		
	B141	EP	0 810 217	A1	JAPAN ENERGY CORPORATION	12-03-1997		
	B142	EP	0 769 947	B1	TANG, Peng Cho et al.	05-02-1997		

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	B143	EP	0 934 931	A2	SUGEN, INC.	08-11-1999		
	B144	EP	1 082 305	A1	SUGEN, INC.	03-14-2001		
	B145	FR	1.398.224		IMPERIAL CHEMICAL INDUSTRIES LIMITED	05-07-1965		X
	B146	FR	1.599.772		INSTITUT PASTEUR	08-28-1970		X
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	B148	GB	809,691		Roy HULL	03-04-1959		
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	B150	JP	62-29570	A	KANEGAFUCHI CHEM KK	02-07-1987		X
	B151	JP	62-39564	A	KANEGAFUCHI CHEM KK	02-20-1987		X
	B152	JP	63-141955	A	KANEGAFUCHI CHEM KK	06-14-1988		X
	B153	JP	5-58894	A	KANEKA CORP	03-09-1993		X
	B154	CA	2,012,634	A1	UNIVERSITY OF BRITISH COLUMBIA	09-20-1991		
	B155	AU	286870		IMPERIAL CHEMICAL INDUSTRIES OF AUSTRALIA AND NEW ZEALAND LIMITED	05-11-1967		

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	B156	ABRAMOVITCH and HEY, "Internuclear cyclisation. Part VIII. Naphth[3:2:1-cd]oxindoles," <u>J. Chem. Soc.</u> 1697-1703 (1954), Strand, London		
	B157	ABRAMOVITCH et al., "A Novel Synthesis of a Cyclic Hydroxamic Acid Involving a Molecular Rearrangement," <u>Chemistry and Industry</u> 44:1871 (1967) 8Laporte Industries Limited, Lancashire		
	B158	Beilstein Reg. No. 236050, Beilstein Reference No. 4-21-00-06355		
	B159	AKBASAK and SUNAR-AKBASAK, "Oncogenes: cause or consequence in the development of glial tumors," <u>J. Neurol. Sci.</u> 111:119-133 (1992) 8 Elsevier Science Publishers		
	B160	ANDREANI et al., "Potential Antitumor Agents. 25[1]. Synthesis and Cytotoxic Activity of 3-(2-Chloro-3-Indolymethylene)1,3-Dihydroindol-2-Ones," <u>Anticancer Research</u> 16:3585-3588 (1996) 8 Elsevier, Paris		
	B161	ANDREANI et al., "Synthesis and cardiotoxic activity of 2-indolinones," <u>Eur. J. Med. Chem.</u> 25:187-190 (1990)		
	B162	ANDREANI et al., "Synthesis and cardiotoxic activity of 2-indolinones bearing pyridyl groups," <u>Eur. J. Med. Chem.</u> 28:653-657 (1993) 8 Elsevier, Paris		
	B163	ANDREANI et al., "Synthesis and cardiotoxic activity of 2-indolinones," <u>Chemical Abstracts</u> , Vol. 113, abstract no. 78106 (1990)		
	B164	ANDREANI et al., "Synthesis and cardiotoxic activity of pyridylmethylene-2-indolinones," <u>Eur. J. Med. Chem.</u> 27:167-170 (1992) 8 Elsevier, Paris		
	B165	ANDREANI et al., "Synthesis and potential coanthracyclic activity of substituted 3-(5-imidazo[2,1-b]thiazolylmethylene)-2-indolinones," <u>Eur. J. Med. Chem.</u> 32:919-924 (1997) 8 Elsevier, Paris		
	B166	ANDREANI et al., "Synthesis of lactams with potential cardiotoxic activity," <u>Eur. J. Med. Chem.</u> 28:825-829 (1993)		
	B167	ANDREANI et al., "In Vivo Cardiotoxic Activity of Pyridylmethylene-2-indolinones," <u>Arzneimittel-Forschung Drug Research</u> 48:727-729 (1998) 8		

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				<b>Prior Appl. Filing Date</b>	February 15, 2002
				<b>First Named Inventor</b>	Huiping GUAN et al.
				<b>Group Art Unit</b>	Unassigned
				<b>Examiner Name</b>	Unassigned
<b>Date Submitted:</b>				<b>Attorney Docket Number</b>	034536-0188
(use as many sheets as necessary)					
Sheet	11	of	30		

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	B168	ARTEAGA et al., "Blockade of the Type I Somatostatin Receptor Inhibits Growth of Human Breast Cancer Cells in Athymic Mice," <u>J. Clin. Invest.</u> 84:1418-1423 (1989) copyright The American Society for Clinical Investigation, Inc.		
	B169	ARVIDSSON et al., "Tyr-716 in the Platelet-Derived Growth Factor $\beta$ -Receptor Kinase Insert is Involved in GRB2 Binding and Ras Activation," <u>Molecular and Cellular Biology</u> 14:6715-6726 (1994) © The American Society for Microbiology		
	B170	AUTREY and TAHK, "The Synthesis and Stereochemistry of Some Isatylideneacetic Acid Derivatives," <u>Tetrahedron</u> 23:901-917 (1967) © Pergamon Press		
	B171	BAHNER and BROTHERTON, "6-Dimethylaminochrysene and Other Analogs of 4-(4-Dimethylamino)stilbene," <u>J. Med. Chem.</u> 12:722-723 (1969)		
	B172	BAHNER et al., "Benzylideneindenes with Oxygen Attached to the Indene Ring," <u>J. Med. Chem.</u> 12:721-722 (1969)		
	B173	BAMFIELD et al., "Diels-Alder Reactions of Oxindolylideneacetone," <u>J. Chem. Soc. (C)</u> 1028-1030 (1966) ©		
	B174	BARBIER, et al., "Synthesis of Isobrassilexin, A Biologically Active Isomer of Brassilexin, a Cruciferae Phytoalexin," <u>Synthetic Communications</u> 23(22):3109-3117 (1993) © Marcel Dekker, Inc.		
	B175	BASERGA, "Oncogenes and the Strategy of Growth Factors," <u>Cell</u> 79:927-930 (1994) © Cell Press		
	B176	BASERGA, "The Insulin-like Growth Factor I Receptor: A Key to Tumor Growth?" <u>Cancer Research</u> 55:249-252 (1995)		

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	B177	Beilstein Reg. No. 233511 (1997)		
	B178	Beilstein Reg. No. 235647 (1997)		
	B179	Beilstein Reg. No. 252929 (1998)		
	B180	BENZIES, et al., "2-Formyl-3-Methoxymethylindole, 3-Ethoxymethyl-2-Formylindoline and 2-Formyl-3-Methylindole," <u>Synthetic Communications</u> : 16(14), 1799-1807 (1986) § Merck Dekker, Inc.		
	B181	BLAKE and JAUQUES, "Anisotropic Effects in $\alpha$ -Substituted Methoxystylenes," <u>J. Chem. Soc. Perkin II</u> : 1660-1663 (1973) § Pergamon, Oxford		
	B182	BOLEN et al., "The Src family of tyrosine protein kinases in hemopoietic signal transduction," <u>FASEB J.</u> 6:3403-3409 (1992)		
	B183	BOLEN, "Nonreceptor tyrosine protein kinases," <u>Oncogene</u> 8:2025-2031 (1993) copyright MacMillan Press Ltd.		
	B184	BONNER et al., "Structure and Biological Activity of Human Homologs of the <i>ras/mil</i> Oncogene," <u>Molecular and Cellular Biology</u> 5:1400-1407 (1985) § The American Society for Microbiology		
	B185	BORSCHKE et al., "Über vielkernige kondensierte Systeme mit heterocyclischen Ringen. XIII," <u>Liebigs Ann. Chem.</u> 550:160-174 (1941)		

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		<b>Group Art Unit</b>	Unassigned
		<b>Examiner Name</b>	Unassigned
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	B187	CANCE et al., "Novel Protein Kinases Expressed in Human Breast Cancer," <u>Int. J. Cancer</u> 54:571-577 (1993) & Wiley-Liss, Inc.	
	B188	CANOIRA and RODRIGUEZ, "Synthesis of Oxindole Derivatives from N-Alkenyl-o-Chloroanilides with Zero-Valent Nickel Complex," <u>J. Heterocyclic Chem.</u> 22:1511-1518 (1985)	
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	B190	CHAO, "Growth Factor Signaling: Where Is the Specificity?" <u>Cell</u> 68:995-997 (1992) copyright Cell Press	
	B191	CHATTEN et al., "Substituted Oxindoles. Part VI. Polarographic Reduction of Substituted <i>trans</i> -3-Benzylideneindol-2(3H)-ones," <u>J. Chem. Soc. Perkin II</u> : 469-473 (1973)	
	B192	CHATTERJEE, et al., "Acylation of Indoles by Duff Reaction and Vilsmeier-Haack Formylation and Conformation of <i>N</i> -Formylindoles," <u>J. Org. Chem.</u> , 38:4002-4004 & The American Chemical Society	
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	B194	CLAESSON-WELSH, "Signal Transduction by the PDGF Receptors," <u>Progress in Growth Factor Research</u> 5:37-54 (1994) & Elsevier Science Ltd.	

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	B195	CODA et al., "(Z)- and (E)-Arylidene-1,3-dihydroindol-2-ones: Configuration, Conformation and Infrared Carbonyl Stretching Frequencies," <u>J. Chem. Soc. Perkin Trans. II</u> : 615-619 (1984)	
	B196	CODA et al., "3-(4-methylbenzylidene)-1,3-dihydroindol-2-one," <u>Journal of the Chemical Society, Perkin Transactions 2</u> 4:615-620 (1984) DATABASE CROSSFIRE, Beilstein Reference No. 6-21	
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	B200	DATI et al., "Inhibition of c-erbB-2 oncogene expression by estrogens in human breast cancer cells," <u>Oncogene</u> 5:1001-1006 (1990)	
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	B202	DE VRIES et al., "The fms-Like Tyrosine Kinase, a Receptor for Vascular Endothelial Growth Factor," <u>Science</u> 255:989-991 (1992)	
	B203	DECKER and LOHMANN-MATTHES, "A quick and simple method for the quantitation of lactate dehydrogenase release in measurements of cellular cytotoxicity and tumor necrosis factor (TNF) activity," <u>J. Immunol. Methods</u> 15:61-69 (1988) copyright Elsevier	

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Prior Application Number	10/076,140
Prior Appl. Filing Date	February 15, 2002
First Named Inventor	Huiping GUAN et al.
Group Art Unit	Unassigned
Examiner Name	Unassigned
Attorney Docket Number	034536-0188

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	B204	DECODTS et al., "Suicide inhibitors of proteases. Lack of activity of halomethyl derivatives of some aromatic lactams," <u>Eur. J. Med. Chem</u> 18: 107-111 (1983)	
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	B206	DICKSON et al., "13. Tyrosine kinase receptor - nuclear protooncogene interactions in breast cancer," <u>Cancer Treatment Res.</u> 61:249-273 (1992) 8 Kluwer Academic Publishers	
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	B220	GAZIT et al., "Tyroshostins. 2. Heterocyclic and $\alpha$ -Substituted Benzylidenemalononitrile Tyroshostins as Potent Inhibitors of EGF Receptor and ErbB2/neu Tyrosine Kinases," <u>J. Med. Chem.</u> 34:1896-1907 (1991) copyright Am. Chem. Soc.		
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	B231	HOWARD, Harry R., "Lactam Derivatives," U.S. Provisional Patent Application Number 60/015134	
	B232	HOWARD et al., "Synthesis and aldose reductase inhibitory activity of substituted 2(1H)-benzimidazolone- and oxindole-1-acetic acids," <u>Eur. J. Med. Chem.</u> 27:779-789 (1992) & Elsevier, Paris	
	B233	HU et al., "Interaction of Phosphatidylinositol 3-Kinase-Associated p85 with Epidermal Growth Factor and Platelet-Derived Growth Factor Receptors," <u>Molecular and Cellular Biology</u> 12:981-990 (1992) copyright Am. Soc. Microbiol.	
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	B239	KATRITZKY et al., "Color and Constitution. Part 8[1]. Some Novel Dyes containing Indoxyl Residues," <u>J. Heterocyclic Chem.</u> 25:1287-1292 (1988)	

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	B240	KAZLAUSKAS et al., "The 64-kDa protein that associates with the platelet-derived growth factor receptor $\beta$ subunit via Tyr-1009 is the SH2-containing phosphotyrosine phosphatase Syp," <u>Proc. Natl. Acad. Sci. USA</u> 90:6939-6942 (1993)		
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	B243	KIKUMOTO et al., "The Reactions of Oxindoles and Isatin with Nitrobenzyl Chlorides," <u>Tetrahedron</u> 22: 3337-3343 (1966) $\S$ Pergamon Press Ltd.		
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	B249	KOHLER and MILSTEIN, "Continuous cultures of fused cells secreting antibody of predefined specificity," <u>Nature</u> 256:495-497 (1975)		
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	B257	LAROCK and BABU, "Synthesis of Nitrogen Heterocycles via Palladium-catalyzed Intramolecular Cyclization," <u>Tetrahedron Letters</u> 28:5291-5294 (1987) copyright Pergamon Journals Ltd.		

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	B258	LEE and DONOGHUE, "Intracellular Retention of Membrane-Anchored v-sis Protein Abrogates Autocrine Signal T transduction," <u>J. Cell. Biol.</u> 118:1057-1070 (1992) §The Rockefeller University Press	
	B259	LEVITZKI and GAZIT, "Tyrosine Kinase Inhibition: An Approach to Drug Development," <u>Science</u> 267:1782-1788 (1995)	
	B260	MAASS et al., "Viral Resistance to the Thiazolo-Iso-Indolinones, a New Class of Nonnucleoside Inhibitors of Human Immunodeficiency Virus Type 1 Reverse Transcriptase," <u>Antimicrobial Agents and Chemotherapy</u> 37:2612-2617 (1993) §American Society for Microbiology	
	B261	MACAULAY et al., "Autocrine Function for Insulin-like Growth Factor I in Human Small Cell Lung Cancer Cell Lines and Fresh Tumor Cells," <u>Cancer Research</u> 50:2511-2517 (1990)	
	B262	MARIANI et al., "Inhibition of angiogenesis by FCE 26806, a potent tyrosine kinase inhibitor," <u>Experimental Therapeutics - Proceedings of the American Association for Cancer Research</u> 35:381 at abstract no. 2268 (March 1994)	
	B263	MARTIN-LEON et al., "On the Cyclization to the Elusive Amino-4H-pyran Ring Some New Facts," <u>Liebigs Ann. Chem.</u> 101-104 (1990) copyright VCH Verlagsgesellschaft mbH & VCH	
	B264	MEL'NIKOVA TV et al., "Indole chemistry. XXXVIII. Cleavage of a carbon-carbon bond during the reaction of 2-aminoindoles with difunctional compounds," <u>Chemical Abstracts</u> 80 (1974) Abstract No. 003413	
	B265	MILLAUER et al., "High Affinity VEGF Binding and Developmental Expression Suggest Flk-1 as a Major Regulator of Vasculogenesis and Angiogenesis," <u>Cell</u> 72:835-846 (1993) § Cell Press	
	B266	MOHAMMADI et al., "Structures of the Tyrosine Kinase Domain of Fibroblast Growth Factor Receptor in Complex with Inhibitors," <u>Science</u> 276:955-960 (1997) § American Association for the Advancement of Science	

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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>				<b>Prior Application Number</b>	10/076,140
				<b>Prior Appl. Filing Date</b>	February 15, 2002
				<b>First Named Inventor</b>	Huiping GUAN et al.
				<b>Group Art Unit</b>	Unassigned
				<b>Examiner Name</b>	Unassigned
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	B267	MORETO et al., "Study of the Laxative Properties of the Disodium Salt of the Sulfuric Diester of 3,3 Bis-(4-Hydroxyphenyl)-7-Methyl-2-Indolinone (DAN-603) in the Rat," <u>European Journal of Pharmacology</u> 36:221-226 (1976) North-Holland Publishing Company		
	B268	MORETO et al., "3,3-Bis-(4-Hydroxyphenyl)-7-Methyl-2-Indolinone (BHMI), the Active Metabolite of the Laxative Sulisatin," <u>Arzneimittel-Forschung Drug Research</u> 29:1561-1564 (1979)		
	B269	MORRISON et al., "Signal Transduction From Membrane to Cytoplasm: Growth Factors and Membrane-Bound Oncogene Products Increase Raf-1 Phosphorylation and Associated Protein Kinase Activity," <u>Proc. Natl. Acad. Sci. USA</u> 85:8855-8859 (1988)		
	B270	MOSMANN, "Rapid Colorimetric Assay for Cellular Growth and Survival: Application to Proliferation and Cytotoxicity Assays," <u>J. Immunol. Methods</u> 65:55-63 (1983) copyright Elsevier Publishers B.V.		
	B271	NEBER and RÖCKER, "On the action of benzaldehydes on the free o-aminophenylacetic acid (II)," <u>Chem. Ber.</u> 56:1710-1716 (1923) (GERMAN AND ENGLISH TRANSLATION)		
	B272	NISHIMURA et al., "Two Signaling Molecules Share a Phosphotyrosine-Containing Binding Site in the Platelet-Derived Growth Factor Receptor," <u>Molecular and Cellular Biology</u> 13:6889-6896 (1993)		
	B273	NODIFF et al., "Antimalarial Phenanthrene Amino Alcohols. 1. Fluorine-Containing 3- and 6-Substituted 9-Phenanthrenemethanols," <u>J. Med. Chem.</u> 14:921-925 (1971)		
	B274	NODIFF et al., "Antimalarial Phenanthrene Amino Alcohols. 3. Halogen-containing 9-phenanthrenemethanols," <u>Chemical Abstracts</u> , Vol. 83, abstract no. 188214 (1975)		
	B275	OSBORNE et al., "Effect of Estrogens and Antiestrogens on Growth of Human Breast Cancer Cells in Athymic Nude Mice," <u>Cancer Research</u> 45:584-590 (1985)		

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	B276	O'SULLIVAN and ROTHERY, "The Preparation and Possible Clinical Significance of 4'-Dialkylaminoisoidindogenides," <i>Clinica Chimica Acta</i> 62:181-182 (1975) sElsevier Scientific Publishing Company		
	B277	OZZELLO and SORDAT, "Behavior of Tumors Produced by Transplantation of Human Mammary Cell Lines in Athymic Nude Mice," <i>Eur. J. Cancer</i> 16:553-559 (1980)		
	B278	PAVLENKO et al., "Introduction of aminomethyl groups into heterocyclic CH-acid molecules," <i>Dopov. Akad. Nauk Ukr Rrsr. Ser. B: Geol., Khim. Biol. Nauki</i> 7:64-66 (1980) We should add thqat we are Sub. Abstract		
	B279	PERKIN et al., "Harmine and Harmaline. Part II. The Synthesis of isoHarman," <i>J. Chem. Soc.</i> 103:1973-1985 (1913)		
	B280	PLATE et al., "Vascular endothelial growth factor is potential tumor angiogenesis factor in human <u>gliomas in vivo</u> ," <i>Nature</i> 359:845-848 (1992)		
	B281	PLOWMAN et al., "Receptor Tyrosine Kinases as Targets for Drug Intervention," <i>DN&amp;P</i> 7:334-339 (1994)		
	B282	QUALLICH et al., A General Oxindole Synthesis," <i>J. Synthetic Organic Chemistry</i> : 51-51 (1993)		
	B283	QUINN et al., "Fetal liver kinase 1 is a receptor for vascular endothelial growth factor and is selectively expressed in vascular endothelium," <i>Proc. Natl. Acad. Sci. USA</i> 90:7533-7537 (1993)		
	B284	ROZAKIS-ADCOCK et al., "Association of the Shc and Grb2/Sem5 SH2-containing proteins is implicated in activation of the Ras pathway by tyrosine kinases," <i>Nature</i> 360:689-692 (1992)		

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	B285	RUVEDA and GONZALEZ, "Geometric isomerism in benzyldeneoxindoles," <u>Spectrochimica Acta</u> 26A:1275-1277 (1970)		
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	B287	SAINSBURY et al., "Electrochemical Oxidation of Aromatic Ethers. Part 5. <sup>1</sup> Further Studies of the Coupling Reactions of Alkoxyated Alkyl- and Aryl-amides," <u>J.C.S. Perkin I</u> 108-114		
	B288	SAITO and STREULI, "Molecular Characterization of Protein Tyrosine Phosphatases," <u>Cell Growth &amp; Differentiation</u> 2:59-65 (1991) 8Molecular Biolody Journal of the American Association for Cancer Research		
	B289	SANDBERG-NORDQVIST et al., "Characterization of Insulin-Like Growth Factor 1 in Human Primary Brain Tumors," <u>Cancer Research</u> 53:2475-2478 (1993)		
	B290	SCHINDLER et al., "Über Dibenz[b,f]-azocin-Derivate," <u>Helvetica Chimica Acta</u> 49:985-989 (1966)		
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	B292	SCHUCHTER et al., "Successful Treatment of Murine Melanoma with Bryostatins 1," <u>Cancer Research</u> 51:682-687 (1991)		
	B293	SEIBERT et al., "Clonal Variation of MCF-7 Breast Cancer Cells in <u>Vitro</u> and in Athymic Nude Mice," <u>Cancer Research</u> 43:2223-2234 (1983)		

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Prior Appl. Filing Date	February 15, 2002
First Named Inventor	Huiping GUAN et al.
Group Art Unit	Unassigned
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	B294	SHAFIE and GRANTHAM, "Role of Hormones in the Growth and Regression of Human Breast Cancer Cells (MCF-7) Transplanted Into Athymic Nude Mice," <u>J. Natl. Cancer Institute</u> 67:51-56 (1981)	
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	B296	SHIRAIISHI et al., "Specific inhibitors of Tyrosine-Specific Protein Kinase, Synthetic 4-Hydroxycinnamide Derivatives," <u>Biochemical and Biophysical Research Communications</u> 147:322-328 (1987) Academic Press	
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	B298	SHWEIKI et al., "Vascular endothelial growth factor induced by hypoxia may mediate hypoxia-initiated angiogenesis," <u>Nature</u> 359:843-845 (1992)	
	B299	SINGH et al., "Indolinone Derivatives as Potential Antimicrobial Agents," <u>Zentralbl. Mikrobiol.</u> 144:105-109 (1989) copyright VEB Gustav Fischer Verlag Jena	
	B300	SINGH et al., "Synthesis and Anticonvulsant Activity of New 1-Substituted 1'-Methyl-3-Chloro-2-Oxospiro (Azetidin-3', 4-Indol-2' Ones)," <u>Bollettino Chimico Farmaceutico</u> 133:76-79 (1994)	
	B301	SKEHAN et al., "New Colorimetric Cytotoxicity Assay for Anticancer-Drug Screening," <u>J. Natl. Cancer Inst.</u> 82:1107-1112 (1990)	
	B302	SLAMON et al., "Studies of the HER-2/neu Proto-oncogene in Human Breast and Ovarian Cancer," <u>Science</u> 244:707-712 (1989)	

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	B303	SOLDI et al., "Platelet-Activating Factor (PAF) Induces the Early Tyrosine Phosphorylation of Focal Adhesion Kinase (p125 <sup>FAK</sup> ) in Human Endothelial Cells," <u>Oncogene</u> 13:515-525 (1996) copyright Stockton Press		
	B304	SONGYANG et al., "SH2 Domains Recognize Specific Phosphopeptide Sequences," <u>Cell</u> 72:767-778 (1993) 8 Cell Press		
	B305	SONGYANG et al., "Specific Motifs Recognized by the SH2 Domains of Csk, 3BP2, fps/fes, GRB-2, HCP, SHC, Syk and Vav," <u>Molecular and Cellular Biology</u> 14:2777-2785 (1994) 8 American Society for Microbiology		
	B306	SPADA, et al., "Small molecule inhibitors of tyrosine kinase activity," <u>Expert Opinion on Therapeutic Patents</u> 5:805-817 (1995) 8Ashley Publications		
	B307	STETINOVA et al., "Stereochemistry and Photoisomerisation of Furfurylideneoxindoles," <u>Collection Czechoslov. Chem. Commun.</u> 42:2201-2206 (1977)		
	B308	STOLLE, Beilstein Reg. No. 273650, <u>J. Prakt. Chem.</u> , Vol. 2, page 128 (1930)		
	B309	STOLLE, Beilstein Reg. No. 305045, <u>J. Prakt. Chem.</u> , Vol. 2, page 128 (1930)		
	B310	SUMPTER and MILLER, "Chapter IV – Oxindole," in <u>Heterocyclic Compounds With Indole and Carbazole Systems</u> , 8 Interscience Publishers, Inc., New York, pp. 134-153 (1954)		
	B311	SUN et al., "Design, Synthesis, and Evaluations of Substituted 3-[(3- or 4-Carboxyethylpyrrol-2-yl)methylidene]indolin-2-ones as Inhibitors of VEGF, FGF, and PDGF Receptor Tyrosine Kinases," <u>Journal of Medicinal Chemistry</u> 42: 5120-5130 (1999) 8American Chemical Society		

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	B313	SUPERTI-FURGA et al., "A functional screen in yeast for regulators and antagonizers of heterologous protein tyrosine kinases," <u>Nature Biotech.</u> 14:600-605 (1996)		
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	B317	TAKANO et al., "Inhibition of angiogenesis by a novel diaminoanthraquinone that inhibits Protein Kinase C," <u>Mol. Bio. Cell</u> 4:358A at abstract no. 2076 (1993)		
	B318	TERRETT et al., "Combinatorial Synthesis - The Design of Compound Libraries and their Application to Drug Discovery," <u>Tetrahedron</u> 51(30):8135-8173 (1995) copyright Pergamon! all even pages missing!		
	B319	THIO et al., "The Interconversion of 2-(2-Aminophenyl)-3-piperolidinone and 3-(2-piperidylmethyl)-2-indolinone: A Reversible N = N' Transacylation," <u>Notes</u> (1971) 479-482		
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Prior Application Number	10/076,140
Prior Appl. Filing Date	February 15, 2002
First Named Inventor	Huiping GUAN et al.
Group Art Unit	Unassigned
Examiner Name	Unassigned
Attorney Docket Number	034536-0188

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	B321	TORP et al., "Expression of the Epidermal Growth Factor Receptor Gene in Human Brain Metastases," <u>APMIS</u> 100:713-719 (1992)	
	B322	TRAXLER, "Protein tyrosine kinase inhibitors in cancer treatment," <u>Expert Opinion on Therapeutic Patents</u> 7(6):571-588 (1997) & Ashley Publications Ltd.	
	B323	TSAL et al., "The Effect of 3,3-Di-Pyridyl Methyl-1-Phenyl-2-Indolinone on the Nerve Terminal Currents of Mouse Skeletal Muscles," <u>Neuropharmacology</u> 31:943-947 (1992) & Pergamon Press	
	B324	TUZI et al., "Expression of growth factor receptors in human brain tumours," <u>Br. J. Cancer</u> 63:227-233 (1991)	
	B325	TWAMLEY-STEIN et al., "The Src family tyrosine kinases are required for platelet-derived growth factor-mediated signal transduction in NIH 3T3 cells," <u>Proc. Natl. Acad. Sci. USA</u> 90:7696-7700 (1993)	
	B326	ULLRICH and SCHLESSINGER, "Signal Transduction by Receptors with Tyrosine Kinase Activity," <u>Cell</u> 61:203-212 (1990) copyright Cell Press	
	B327	VAISMAN et al., "Characterization of the Receptors for Vascular Endothelial Growth Factor," <u>J. Biol. Chem.</u> 265:19461-19466 (1990) & The American Society for Biochemistry and Molecular Biology	
	B328	VARMA and GUPTA, "Nucleophilic Reactions of 2-Methyl-3-(4'-carbomethoxyphenyl)-4-quinazolinones with 2-Indolinones," <u>J. Indian Chem. Soc.</u> 66:804-805 (1989) & The Indian Chemical Society	
	B329	VOLLER et al., "Ch. 45 - Enzyme-Linked Immunosorbent Assay," in <u>Manual of Clinical Immunology</u> , 2 <sup>nd</sup> edition, Rose and Friedman editors, American Society of Microbiology, Washington, D.C., pp. 359-371 (1980); @ American Society for Microbiology	

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	B330	WAHL et al., "3-benzilidene-5-methyl-1,3-dihydroindol-2-one," <u>Ann. Chim.</u> 350 (1926), DATABASE CROSSFIRE, Beilstein Reference No. 2-21-00-00290	
	B331	WAHL et al., "Chimie Organique - Sur les iso-indogénides," <u>C.R. Hebd. Seances Acad. Sci.</u> 149:132-134 (1909)	
	B332	WAHL, Beilstein Reg. No. 191439, <u>Bull. Soc. Chim. Fr.</u> , page 1038 (1909)	
	B333	WAHL, Beilstein Reg. No. 231732, <u>Bull. Soc. Chim. Fr.</u> , pages 1035-1038 (1909)	
	B334	WALKER, "Synthesis of a $\alpha$ -(p-Aminophenyl)- and $\alpha$ -(p-Chlorophenyl)- $\beta$ -aryl-propionitriles by Catalytic Reduction of Stilbenenitriles," <u>J. Med. Chem.</u> 8:583-588 (1965)	
	B335	WALKER et al., "Synthesis of New 3-(Pyridylmethylene)-, 3-(Pyridylmethyl)-, 3-(Piperidylmethyl)-, and 3-( $\beta$ -Alkylaminoethyl)-2-indolinones. The Reduction of Isoindogénides, Nitro Compounds, and Pyridines in a Series of 2-Indolinones," <u>J. Med. Chem.</u> 8:626-637 (1965)	
	B336	WARRI et al., "Estrogen Suppression of erbB2 Expression is Associated with Increased Growth Rate of ZR-75-1 Human Breast Cancer Cells <i>In Vitro</i> and in Nude Mice," <u>Int. J. Cancer</u> 49:618-623 (1991) & Wiley-Leiss, Inc.	
	B337	WEIDNER et al., "Tumor Angiogenesis and Metastasis -- Correlation in Invasive Breast Carcinoma," <u>New England J. Medicine</u> 324:1-7 (1991) & Massachusetts Medical Society	
	B338	WINKELMANN et al., "Chemotherapeutically Active Nitro Compounds: 4. 5-Nitroimidazoles (Part I)," <u>Arzneim.-Forsch./Drug Res.</u> 27:2251-2263 (1977)	

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	B339	WRIGHT et al., "Cyclic Hydroxamic Acids Derived from Indole," <u>J. Am. Chem. Soc.</u> 78:221-224 (1956)	
	B340	WRIGHT et al., "Inhibition of Angiogenesis in Vitro and In Ovo With an Inhibitor of Cellular Protein Kinases, MDL 27032," <u>J. Cellular Physiology</u> 152:448-457 (1992)	
	B341	YOUNG and BABBITT, "2-(2-Methyl-3-indolyl)-1,4-benzoquinone, a Reversible Redox Substrate at the Carbon-Paste Electrode in Acidic Aqueous-Ethanol Media," <u>J. Org. Chem.</u> 47:1571-1572 (1982) copyright Am. Chem. Soc.	
	B342	ZAMAN et al., "Tyrosine Kinase Activity of Purified Recombinant Cytoplasmic Domain of Platelet-Derived Growth Factor $\beta$ -Receptor ( $\beta$ -PDGFR) and Discovery of a Novel Inhibitor of Receptor Tyrosine Kinases," <u>Biochemical Pharmacology</u> 57:57-64 (1999) Elsevier Science Inc.	
	B343	ZHANG et al., "Microtubule Effects of Welwistatin, a Cyanobacterial Indolinone that Circumvents Multiple Drug Resistance," <u>Molecular Pharmacology</u> 49:228-234 (1996) The American Society for Pharmacology and Experimental Pharmaceutics	
	B344	ZHUNGIETU et al., "Reaction of Indoles and 2-Ketoindolines With Some Aldehydes," <u>Chemical Abstracts</u> , Vol. 78, abstract no. 111201 (1990)	

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